

Application No. 10/699,393
Response Dated January 18, 2007
Second Reply to Office Action of August 24, 2006

RECEIVED
CENTRAL FAX CENTER
JAN 18 2007

Amendments to the Specification

Please replace the paragraph on page 1, lines 6-7 with the following replacement paragraph:

The present application is a divisional application of, and claims benefit of U.S. Patent Application Serial No. 10/165,422, filed June 7, 2002, entitled "Antithrombotic thrombin variants," now issued as U.S. Pat. No. 6,706,512, which itself claims benefit of priority from U.S. Provisional Application Serial No. 60/297,089, filed June 8, 2001, ~~the disclosure of which is hereby incorporated herein in its entirety by reference.~~

Please replace the paragraph on page 7, lines 17-28 with the following replacement paragraph:

Fig. 1 illustrates the amino acid sequence of the thrombin variant W215A (SEQ ID NO:1). The substitution W215A is in bold. The amino acid numbering system herein is based on the Sadler numbering scheme. The A chain of thrombin is designated with "a" postscripts, as in T1a to R36a, while the B chain commences with I1 and extends to E259.

Fig. 2 illustrates the amino acid sequence of the thrombin variant W215A B-chain (SEQ ID NO:2). The amino acid numbering system herein is based on the Sadler numbering scheme. The B chain commences with I1 and extends to E259.

Fig. 3 illustrates the amino acid sequence of the thrombin variant W215A/E217A (SEQ ID NO:3). The amino acid substitutions W215A and E217A are in bold. The amino acid numbering system herein is based on the Sadler numbering scheme. The A chain of thrombin is designated with "a" postscripts, as in T1a to R36a, while the B chain commences with I1 and extends to E259.

Jan-18-2007 02:59pm From-MORRIS MANNING MARTIN

4043644578

T-271 P.007 F-877

Application No. 10/699,393
Response Dated January 18, 2007
Second Reply to Office Action of August 24, 2006

Fig. 4 illustrates the amino acid sequence of the thrombin variant W215A/E217A (WB) B-chain (SEQ ID NO:4). The amino acid numbering system herein is based on the Sadler numbering scheme. The B chain commences with I1 and extends to E259.